

FIGURE 1

Acetylenes

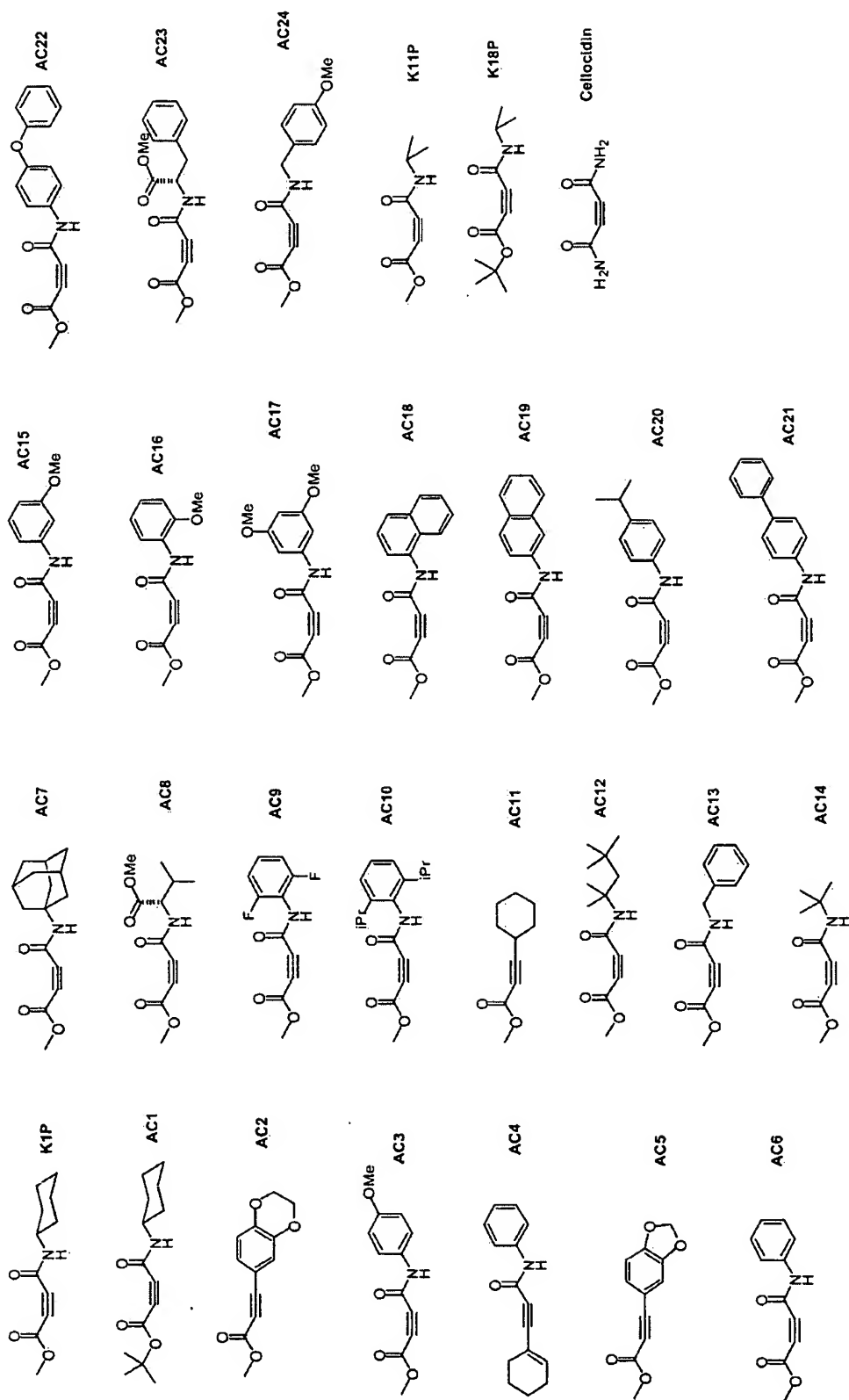


FIGURE 2

Development of planar scaffold molecules

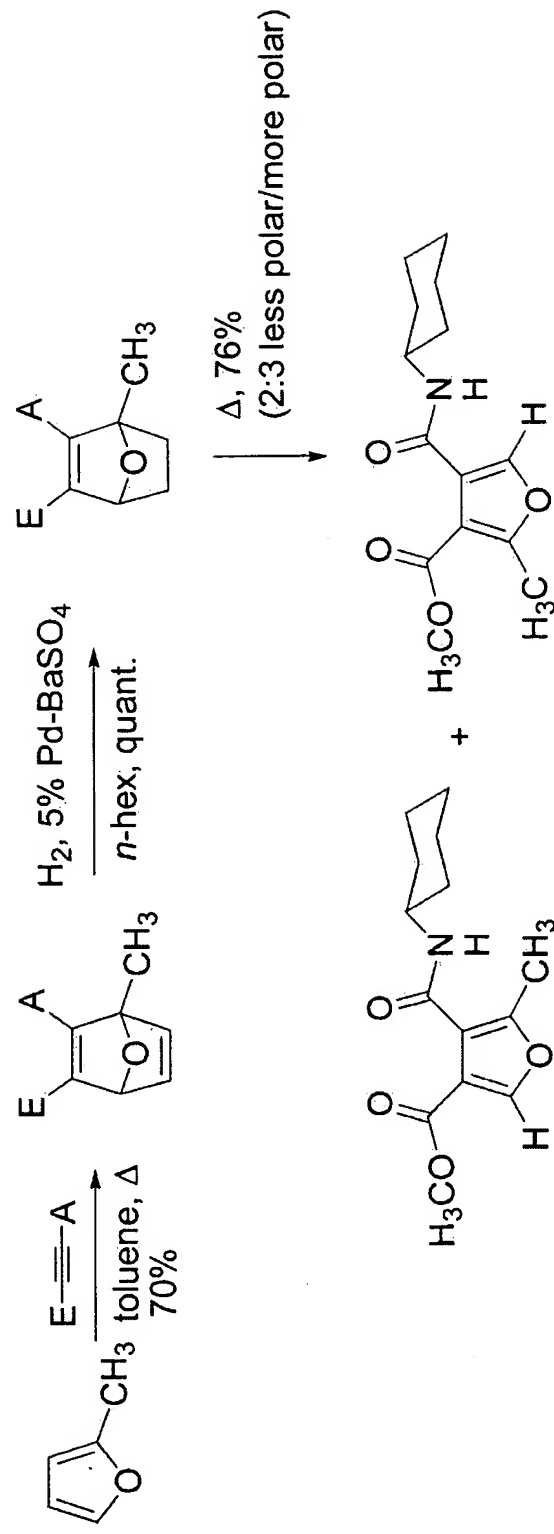


FIGURE 3

Development of planar scaffold molecules, 2

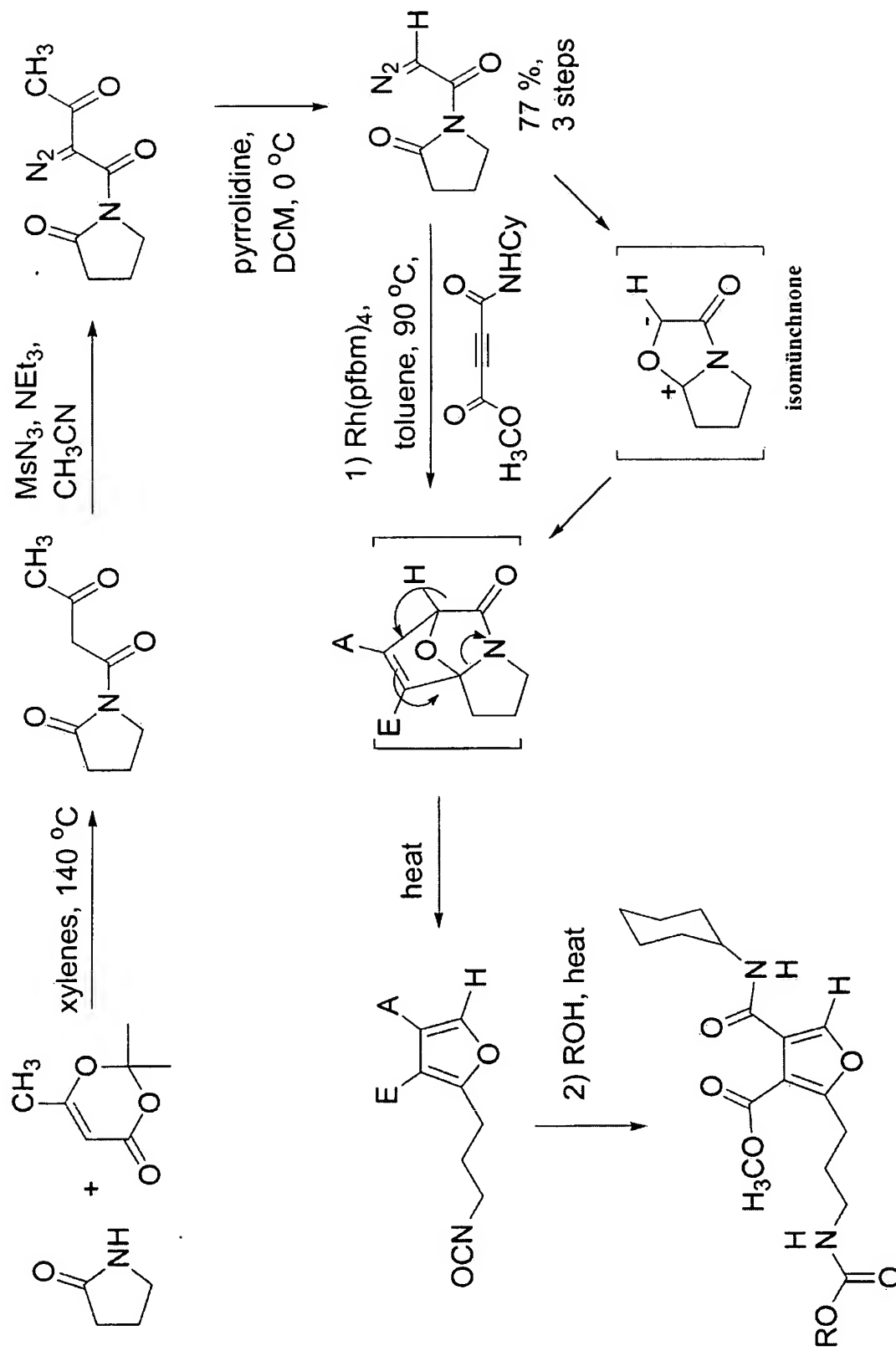


FIGURE 4

Library of Acetylenes

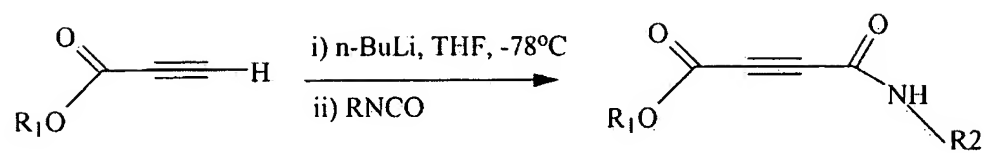
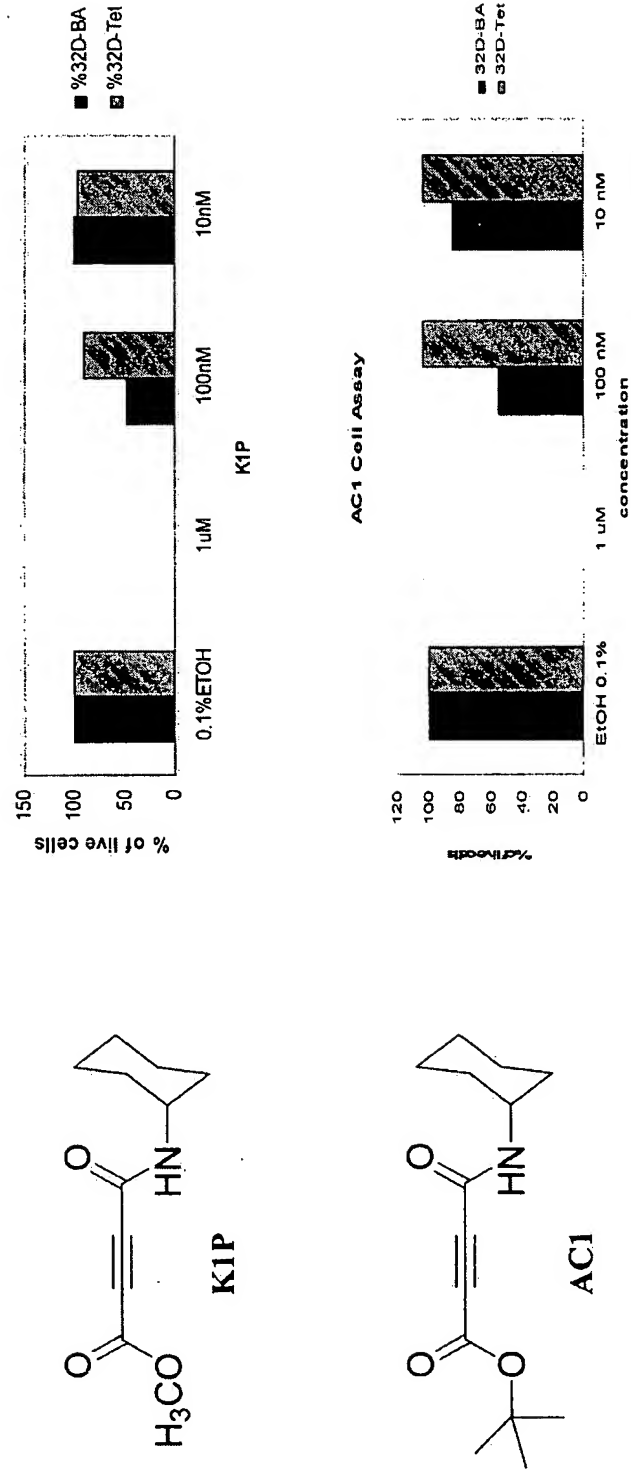


FIGURE 5

1) Investigation of the scaffold left side

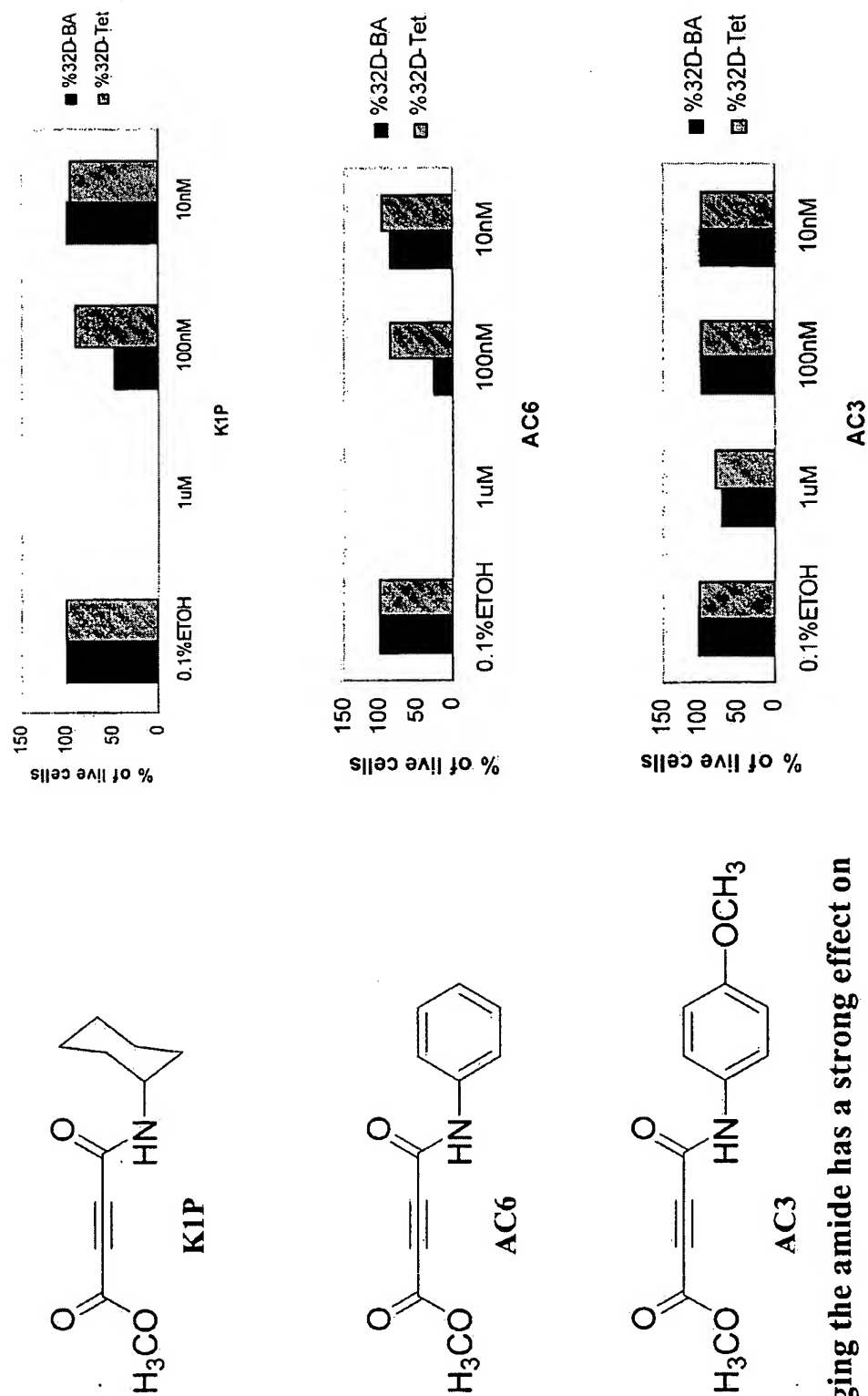


Both molecules exhibited similar activity, despite difference at the ester moiety; optimisation continued with modifications at the amide site

FIGURE 6

Library of Acetylenes

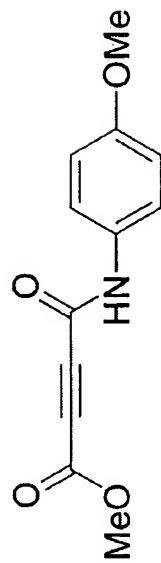
2) Investigation of the scaffold right side



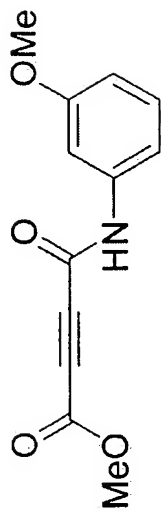
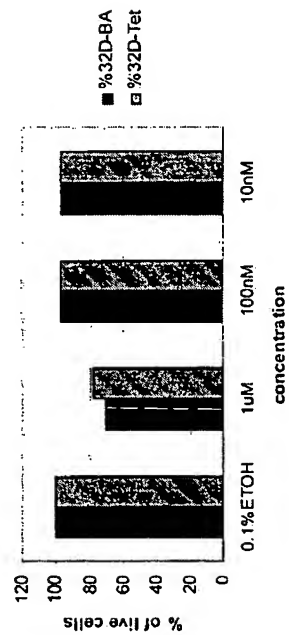
Changing the amide has a strong effect on activity

FIGURE 7

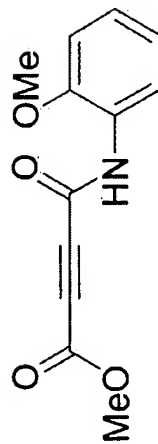
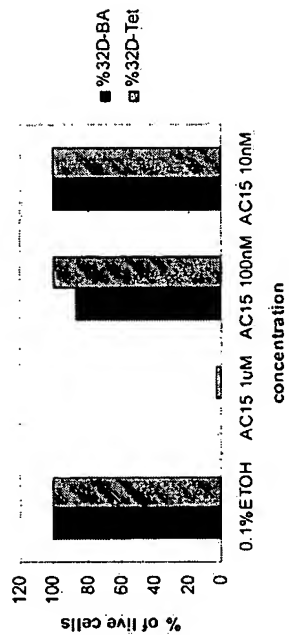
Library of Acetylenes



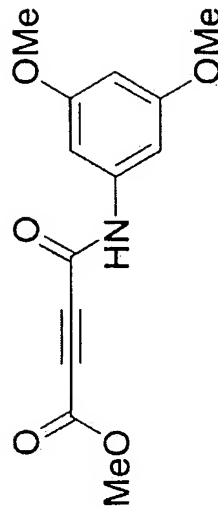
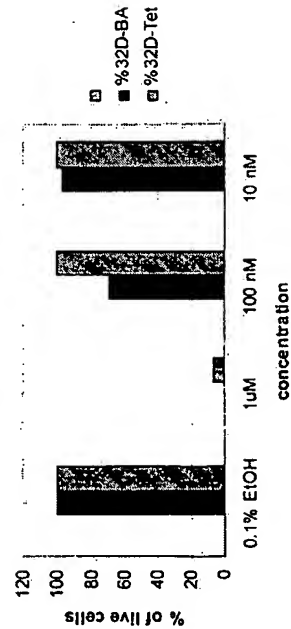
AC3 Cell Assay



AC15 Cell Assay



AC16 Cell Assay



AC17 Cell Assay

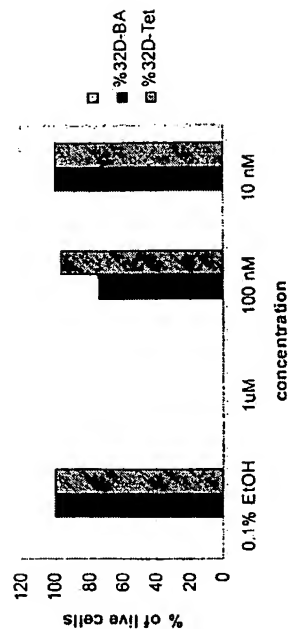
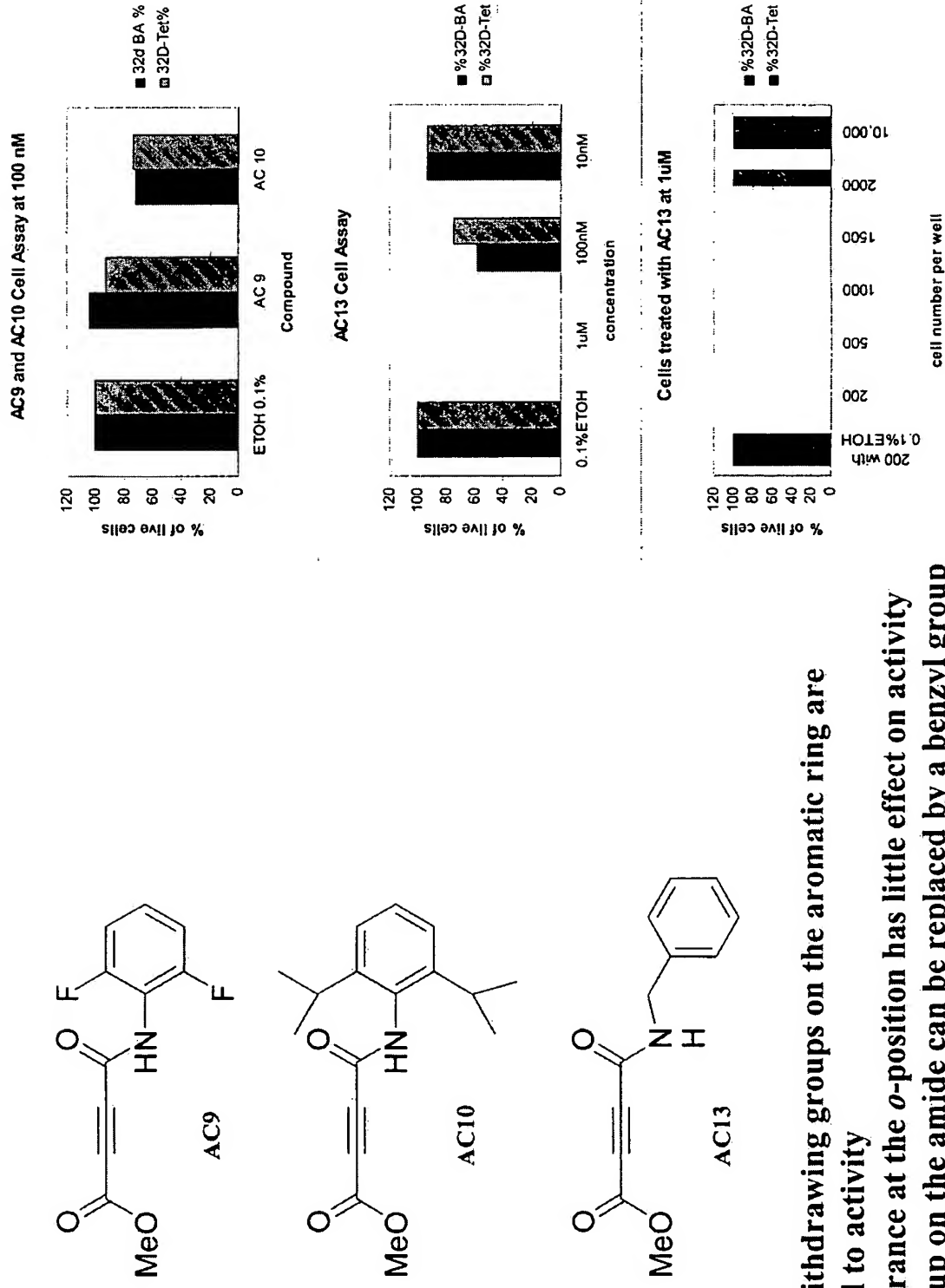


FIGURE 8

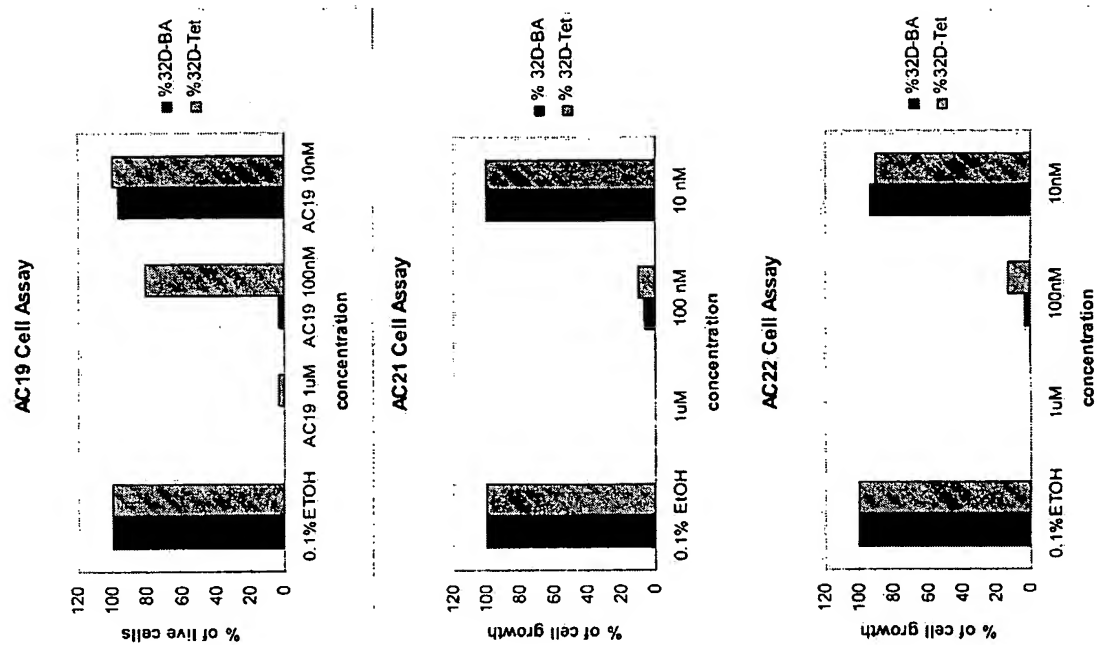
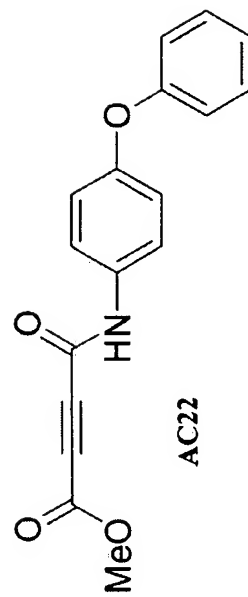
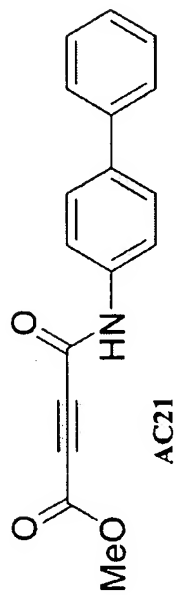
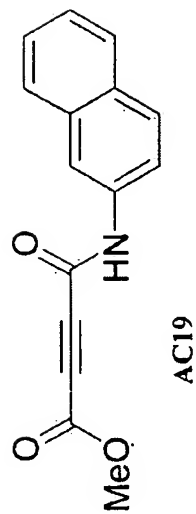
Library of Acetylenes



- Electron withdrawing groups on the aromatic ring are detrimental to activity
- Steric hindrance at the *o*-position has little effect on activity
- Phenyl group on the amide can be replaced by a benzyl group

FIGURE 9

Library of Acetylenes



AC21 and AC22 have lost selectivity displayed by AC19 but not the activity

FIGURE 10

Some acetylenes exhibit synergism with STI-571

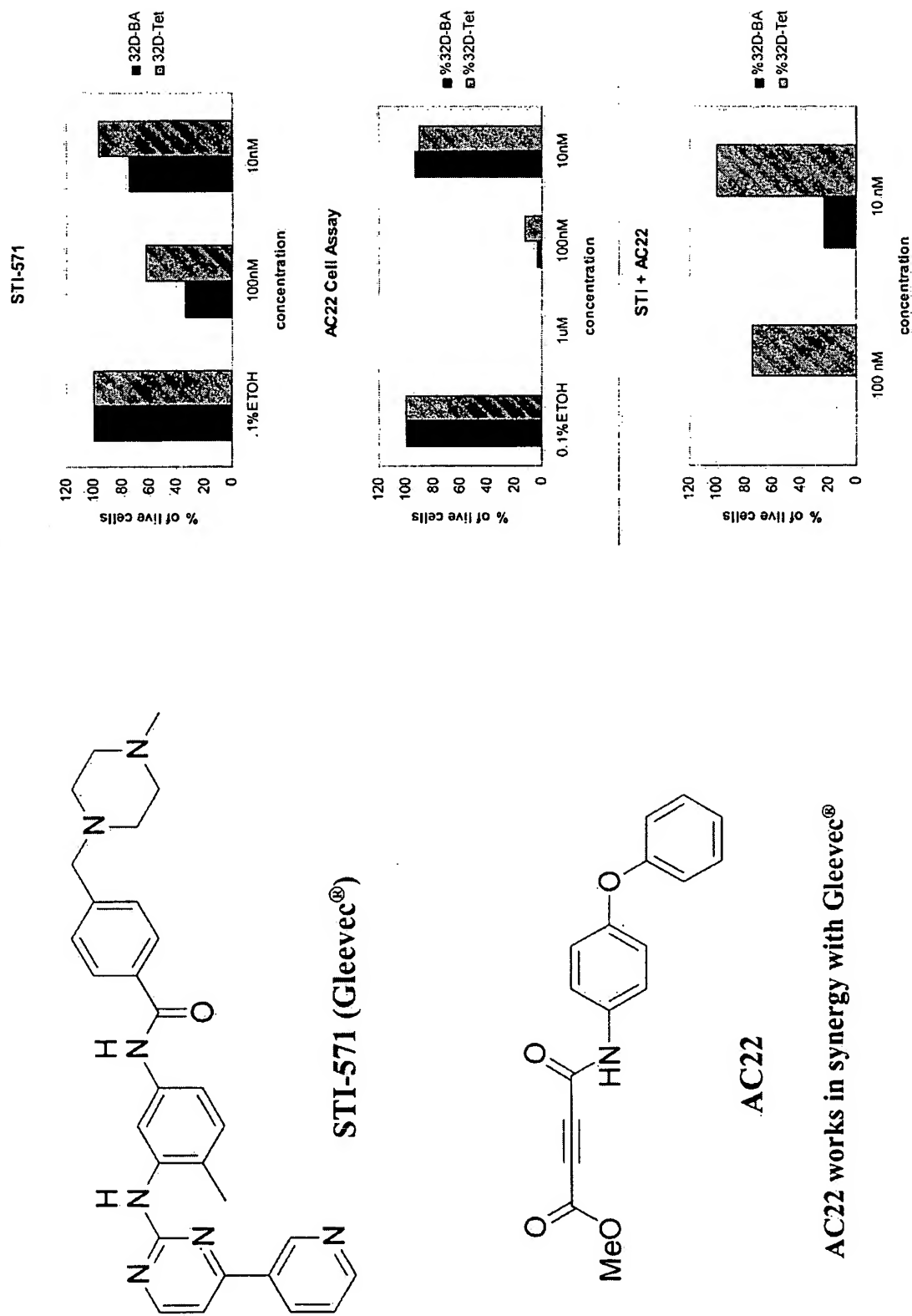


FIGURE 11

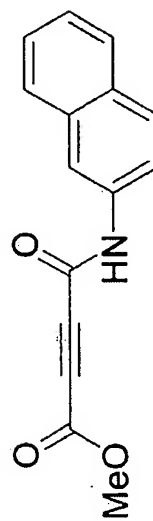
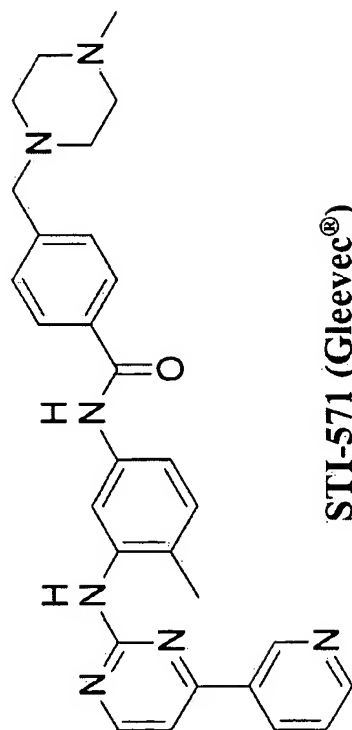
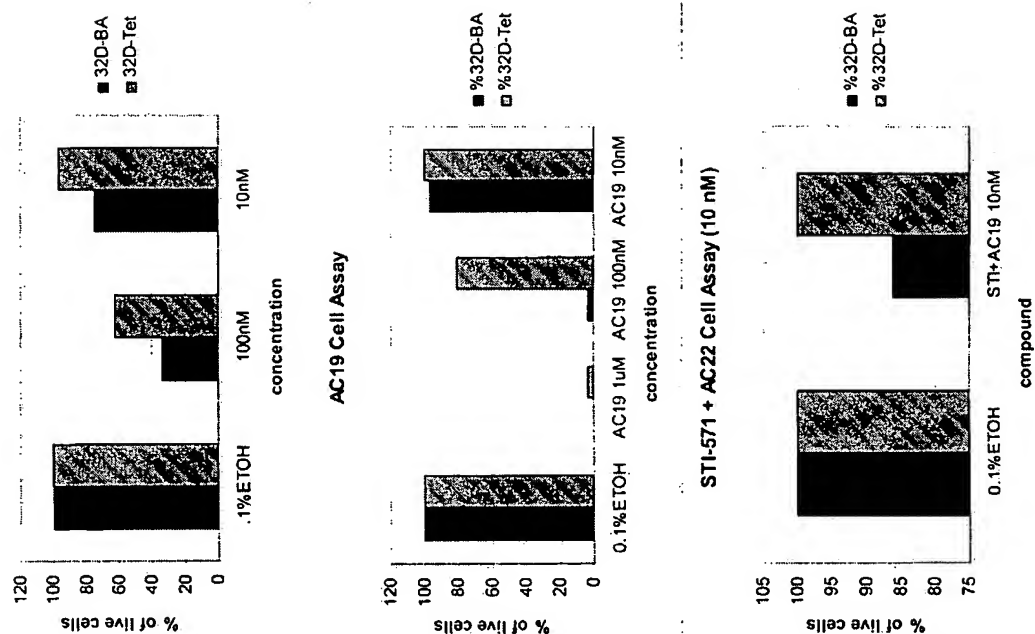


FIGURE 12

Testing of one furan from the second generation furan library

Recently, furans containing a H and a CH₃ were tested and proved to be active

